

- Sub E1* 42. (Amended) A method for generating a culture that is purified or enriched in neural progenitor cells, comprising:
- D2*
- (i) introducing into a pluripotent cell a selectable marker that is differentially expressed in neural progenitor cells compared with its expression in other cells, wherein neural progenitor cells constitute a sub-set of the cells obtainable from the pluripotent cell;
 - (ii) culturing the pluripotent cell *in vitro* to induce differentiation of the pluripotent cell into a neural progenitor cell or into a mixture of cells including neural progenitor cells, or to induce preferential survival, in a mixed culture of cells, of neural progenitor cells; and
 - (iii) selecting for neural progenitor cells according to differential expression of the selectable marker introduced in step (i).

Sub E2 44. (Amended) A method according to Claim 42 wherein the pluripotent cell is selected from embryonic stem (ES) cells, embryonic germ (EG) cells, embryonic carcinoma (EC) cells, a primary culture of fetal cells, a primary culture of post-natal cells, and a primary culture of adult cells.

D3 45. (Amended) A method according to Claim 42 comprising genetically modifying pluripotent cells to delete, mutate, substitute or add genes in order (i) to assay gene function in neural progenitor, and/or (ii) to render selected cells more suitable for transplantation.

46. (Amended) A method according to Claim 42 further comprising:

- (iv) introducing into the pluripotent cell a second selectable marker that is differentially expressed in cells of a selected sub-lineage compared with its expression

FINNEGAN
HENDERSON
FARABOW
CARRETT &
DUNNER LLP

1300 I Street, NW
Washington, DC 20005
202.408.4000
Fax 202.408.4400
www.finnegan.com

in other cells, wherein cells of the selected sub-lineage are formed by differentiation of neural progenitor cells; and

(v) when a culture of neural progenitor cells has been obtained, allowing or inducing differentiation of the cells and selecting for cells that express the second selectable marker.

47. (Twice Amended) A method according to Claim 42 wherein the selectable marker is introduced into the pluripotent cell by targeted integration or random gene trap integration so as to be operatively coupled to a gene that is differentially expressed in neural progenitor cells.

48. (Twice Amended) A method according to Claim 42 wherein the selectable marker is introduced into the pluripotent cell via random integration of a transgene in which the selectable marker is operatively coupled to a gene that is differentially expressed in neural progenitor cells.

49. (Twice Amended) A method according to Claim 42 wherein the pluripotent cell is an ES, EG, or EC cell and the method comprises forming an embryoid body, or otherwise inducing differentiation of the cells.

53. (Amended) A method according to Claim 42 wherein the selectable marker is expressed in cells that express a Sox gene.

REMARKS

Upon entry of this Amendment, claims 42, 44-51, 53, 54, 58, 64, and 65 are pending in this application. Applicants have canceled claims 43, 52, 55-57, and 59-63 without prejudice to their right to prosecute the subject matter of these claims in a divisional application. Applicants have amended the claims to reflect the election made in response to the Office Action dated August 16, 2001. Accordingly, the claims now

FINNEGAN
HENDERSON
FARABOW
GARRETT &
DUNNER LLP

1300 I Street, NW
Washington, DC 20005
202.408.4000
Fax 202.408.4400
www.finnegan.com